

AMENDMENTS**Amendments to the Claims**

1-47. (Canceled)

48. (Currently amended) A method of treating ~~a patient suffering from poisoning or at risk of poisoning by a clostridial toxin~~ in a patient in need thereof, the method comprising the step of supplying administering an effective amount of a toxin-resistant SNAP-25 or a toxin-inhibitory SNAP-25 to the patient; ~~a SNARE (soluble (N-ethylmaleimide-sensitive fusion protein)-attachment protein receptor) to a cell of the patient, wherein the SNARE is resistant to proteolysis by the said clostridial toxin (toxin-resistant SNARE) and/or is capable of inhibiting the clostridial~~ clostridial toxin (toxin-inhibitory SNARE)

wherein the toxin-resistant SNAP-25 is a SNAP-25 resistant to proteolysis by the clostridial toxin;

wherein the toxin-inhibitory SNAP-25 is a SNAP-25 capable of inhibiting the protease activity of the clostridial toxin;

wherein administration of the toxin-resistant SNAP-25 or the toxin-inhibitory SNAP-25 produces a clinically useful or significant reduction in a symptom of poisoning caused by the clostridial toxin in the patient suffering from clostridial toxin poisoning.

49. (Canceled)

50. (Currently amended) A method of preventing poisoning by a clostridial toxin in a patient in need thereof, the method comprising the step of administering an effective amount of a toxin-resistant SNAP-25 or a toxin-inhibitory SNAP-25 to the patient; ~~reversing the inhibition of exocytosis in a cell caused by contact of a clostridial toxin with the said cell, including the step of supplying a SNARE (soluble (N-ethylmaleimide-sensitive fusion protein)-attachment protein receptor) to the said cell not before contact of the said clostridial toxin with the said cell, wherein the SNARE is resistant to proteolysis by the~~

~~said clostridial toxin (toxin-resistant SNARE) and/or is capable of inhibiting the clostridial toxin (toxin-inhibitory SNARE)~~

wherein the toxin-resistant SNAP-25 is a SNAP-25 resistant to proteolysis by the clostridial toxin;

wherein the toxin-inhibitory SNAP-25 is a SNAP-25 capable of inhibiting the protease activity of the clostridial toxin;

wherein administration of the toxin-resistant SNAP-25 or the toxin-inhibitory SNAP-25 produces a clinically useful or significant reduction in a symptom of poisoning caused by the clostridial toxin in the patient at risk of poisoning when exposed to the clostridial toxin.

51-52. (Canceled)

53. (Currently amended) The A-method of either claim 48 or claim 50, as in any one of claims 48-50 wherein the said clostridial toxin is a botulinum toxin type A (BoNT/A).

54. (Currently amended) The A-method as in claim 51 of either claim 48 or claim 50, wherein the said clostridial toxin is botulinum toxin type C1-A (BoNT/A).

55. (Currently amended) The A-method as in claim 52 of either claim 48 or claim 50, wherein the said clostridial toxin is botulinum toxin type E-A (BoNT/A).

56. (Canceled)

57. (Currently amended) The A-method as in claim 54 of either claim 48 or claim 50, wherein the toxin-resistant SNAP-25 or the toxin-inhibitory SNAP-25 comprises a replacement of a said SNARE is a variant of SNAP-25 in which the residue equivalent to residue 197 and/or the residue equivalent to residue 198 of full length SNAP-25 are replaced by a residue other than Q or a residue other than R, respectively.

58. (Currently amended) ~~The A-method as in claim 57~~ of either claim 48 or claim 50, wherein the toxin-resistant SNAP-25 or the toxin-inhibitory SNAP-25 comprises a replacement of a said SNARE is a variant of SNAP-25 in which the residue equivalent to residue 197 and/or the residue equivalent to residue 198 of full length SNAP-25 are replaced by a residue other than Q or a residue other than R, respectively.
59. (Currently amended) ~~The A-method as in~~ of claim 57, wherein the residue equivalent to R198 of full length human SNAP-25 is replaced by a residue other than R, selected from A, T, K, H or W and the residue equivalent to residue Q197 of full length SNAP-25 is Q or is replaced[[.]] by a residue selected from the group consisting of A, K or W.
60. (Currently amended) ~~The A-method as in~~ of claim 58, wherein the residue equivalent to R198 of full length human SNAP-25 is replaced by a residue other than R, selected from the group consisting of A, T, K, H or W and the residue equivalent to residue Q197 of full length SNAP-25 is Q or is replaced, by A, K or W.
61. (Currently amended) ~~The A-method~~ of either claim 48 or claim 50, ~~as in any one of claims 48-50~~ wherein the toxin-resistant SNAP-25 said SNARE that is resistant to proteolysis by the said clostridial toxin is capable of performing substantially the equivalent function of a SNAP-25 endogenously present in the patient to a SNARE present in the cell that is capable of being cleaved in the said cell by the said clostridial toxin.
62. (Currently amended) ~~The A-method~~ of either claim 48 or claim 50, ~~as in claim 51~~ wherein the clostridial toxin poisoning is botulism said SNARE that is resistant to proteolysis by the said clostridial toxin is capable of performing substantially the equivalent function to a SNARE present in the cell that is capable of being cleaved in the said cell by the said clostridial toxin.
- 63-68. (Canceled)

69. (Currently amended) The A-method of either claim 48 or claim 50, ~~as in either of claims 48 or 49~~ wherein the patient is an infant.

70. (Currently amended) The A-method of either claim 48 or claim 50, ~~as in claim 51 wherein the patient is an infant~~ adult.

71-72 (Canceled)

73. (Currently amended) The A-method of either claim 48 or claim 50, ~~as in either of claims 48 or 49~~ further comprising the step of treating the patient with an inhibitor of the said clostridial toxin.

74. (Canceled)

75. (Currently amended) The A-method of claim 73, ~~as in claim 73~~ wherein the clostridial toxin inhibitor is N-acetyl-CRATML-carboximide ~~of the said clostridial toxin is a SNARE that is capable of inhibiting the said clostridial toxin (toxin-inhibitory SNARE) or a recombinant polynucleotide capable of expressing the said toxin-inhibitory SNARE.~~

76-103. (Canceled)